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           Tai, Mei-Sheng
           McCartney, John
     <120> Modified TGF-beta Superfamily Proteins
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     Glu Asn Val Asn Leu Lys Lys Tyr Arg Asn Met Ile Val Lys Ser Cys
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ارارا
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Gly Cys His
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Cys Gly Cys Arg
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Val Val Thr Tyr Lys Phe Lys Tyr Glu Gly Met Ala Val Ser Glu Cys
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Gly Cys Arg
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Gly Cys Ser
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Val Pro Thr Lys Leu Thr Pro Ile Ser Ile Leu Tyr Ile Asp Ala Gly
Asn Asn Val Val Tyr Asn Glu Tyr Glu Glu Met Val Val Glu Ser Cys
Gly Cys Arg
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    Gly Cys Arg
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Gly Cys Arg
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Gly Cys Arg
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Gly Cys Arg
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    Thr Cys Arg
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    Val Tyr His Ile Leu Arg Lys His Ser Ala Lys Arg Cys Gly Cys Ile
                                      25
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15
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    <223> Inhibin Alpha
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    Asp Gly Gly Tyr Ser Phe Lys Tyr Glu Thr Val Pro Asn Leu Leu Thr
    Gln His Cys Ala Cys Ile
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Gln Asn Ile Ile Lys Lys Asp Ile Gln Asn Met Ile Val Glu Glu Cys
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Gly Cys Ser
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Gly Cys Ala
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     Cys Arg
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     Arg Val Leu Leu Glu His His Lys Asp Met Ile Val Glu Glu Cys Gly
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::
     Cys Leu
enè
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     <211> 35
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     <220>
     <223> OP-2
     <400> 26
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     Gly Cys His
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Asn Asn Val Ile Leu Arg Arg Glu Arg Asn Met Val Val Gln Ala Cys
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Gly Cys His
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Asp Ile Ile Asp Leu Thr Lys Tyr Gln Lys Ala Val Ala Lys Glu Cys
Gly Cys His
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Cys Ser
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<223> TGF-Beta2
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Thr Pro Lys Ile Glu Gln Leu Ser Asn Met Ile Val Lys Ser Cys Lys
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Cys Ser
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<213> Gallus gallus
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Cys Ser
<210> 33
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<213> Strongylocentrotus purpuratus
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<223> UNIVIN
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Gly Cys Arg
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<213> Xenopus laevis
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                                                      Met His Val
cgc tca ctg cga gct gcg gcg ccg cac agc ttc gtg gcg ctc tgg gca
                                                                    105
Arg Ser Leu Arg Ala Ala Ala Pro His Ser Phe Val Ala Leu Trp Ala
      5
                         10
ecc etg tte etg etg ege tee gee etg gee gae tte age etg gae aac
                                                                    153
Pro Leu Phe Leu Leu Arg Ser Ala Leu Ala Asp Phe Ser Leu Asp Asn
 20
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gag gtg cac tcg agc ttc atc cac cgg cgc ctc cgc agc cag gag cgg
                                                                    201
Glu Val His Ser Ser Phe Ile His Arg Arg Leu Arg Ser Gln Glu Arg
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cgg gag atg cag cgc gag atc ctc tcc att ttg ggc ttg ccc cac cgc
                                                                    249
Arg Glu Met Gln Arg Glu Ile Leu Ser Ile Leu Gly Leu Pro His Arg
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						ggc Gly										297
						atg Met 90										345
						ccc Pro										393
		_	-	-	_	caa Gln	-	-					_	_	-	441
_	-	-	_		_	aac Asn			_		-					489
						cga Arg										537
						acg Thr 170										585
						gac Asp										633
						ttg Leu										681
						gcc Ala										729
atc Ile	aca Thr	gcc Ala 230	acc Thr	agc Ser	aac Asn	cac His	tgg Trp 235	gtg Val	gtc Val	aat Asn	ccg Pro	cgg Arg 240	cac His	aac Asn	ctg Leu	777
ggc Gly	ctg Leu 245	cag Gln	ctc Leu	tcg Ser	gtg Val	gag Glu 250	acg Thr	ctg Leu	gat Asp	ggg Gly	cag Gln 255	agc Ser	atc Ile	aac Asn	ccc Pro	825
						ggg Gly										873
						aag Lys										921

cgg tcc acg (Arg Ser Thr (,,,				-	_	_		969	
aag aac cag o Lys Asn Gln (310			-				-	-	1017	
agc gac cag a Ser Asp Gln A 325			_		-	_	_		1065	
cga gac ctg o Arg Asp Leu 0 340							Tyr		1113	
gcc tac tac t Ala Tyr Tyr (_		-			_	1161	
aac gcc acc a									1209	
ccg gaa acg o Pro Glu Thr V 390									1257	
atc tcc gtc of Ile Ser Val 1									1305	
tac aga aac a Tyr Arg Asn N 420				_		ctcct	cc		1351	
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cgtttccaga g	gtaattatg a	gcgcctacc	agccago	gcca	cccagcc	gtg g	ggagg	aaggg	1711	
ggcgtggcaa g	gggtgggca c	attggtgtc	tgtgcga	aaag	gaaaatt	gac o	ccgga	agttc	1771	
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<212> PRT

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Leu Asp Asn Glu Val His Ser Ser Phe Ile His Arg Arg Leu Arg Ser 35 40 45

Gln Glu Arg Arg Glu Met Gln Arg Glu Ile Leu Ser Ile Leu Gly Leu
50 55 60

Pro His Arg Pro Arg Pro His Leu Gln Gly Lys His Asn Ser Ala Pro 65 70 75 80

Met Phe Met Leu Asp Leu Tyr Asn Ala Met Ala Val Glu Glu Gly Gly
85 90 95

Gly Pro Gly Gly Gln Gly Phe Ser Tyr Pro Tyr Lys Ala Val Phe Ser 100 105 110

Thr Gln Gly Pro Pro Leu Ala Ser Leu Gln Asp Ser His Phe Leu Thr 115 120 125

Asp Ala Asp Met Val Met Ser Phe Val Asn Leu Val Glu His Asp Lys 130 135 140

Glu Phe Phe His Pro Arg Tyr His His Arg Glu Phe Arg Phe Asp Leu 145 150 155 160

Ser Lys Ile Pro Glu Gly Glu Ala Val Thr Ala Ala Glu Phe Arg Ile 165 170 175

Tyr Lys Asp Tyr Ile Arg Glu Arg Phe Asp Asn Glu Thr Phe Arg Ile 180 185 190

Ser Val Tyr Gln Val Leu Gln Glu His Leu Gly Arg Glu Ser Asp Leu 195 200 205

Phe Leu Leu Asp Ser Arg Thr Leu Trp Ala Ser Glu Glu Gly Trp Leu 210 215 220

Val Phe Asp Ile Thr Ala Thr Ser Asn His Trp Val Val Asn Pro Arg 225 230 235 240

His Asn Leu Gly Leu Gln Leu Ser Val Glu Thr Leu Asp Gly Gln Ser 245 250 255

Ile Asn Pro Lys Leu Ala Gly Leu Ile Gly Arg His Gly Pro Gln Asn 260 265 270

Lys Gln Pro Phe Met Val Ala Phe Phe Lys Ala Thr Glu Val His Phe 275 280 285

Arg Ser Ile Arg Ser Thr Gly Ser Lys Gln Arg Ser Gln Asn Arg Ser

290 295 300

Lys Thr Pro Lys Asn Gln Glu Ala Leu Arg Met Ala Asn Val Ala Glu 305 310 315 320

Asn Ser Ser Ser Asp Gln Arg Gln Ala Cys Lys Lys His Glu Leu Tyr 325 330 335

Val Ser Phe Arg Asp Leu Gly Trp Gln Asp Trp Ile Ile Ala Pro Glu 340 345 350

Gly Tyr Ala Ala Tyr Tyr Cys Glu Gly Glu Cys Ala Phe Pro Leu Asn 355 360 365

Ser Tyr Met Asn Ala Thr Asn His Ala Ile Val Gln Thr Leu Val His 370 380

Phe Ile Asn Pro Glu Thr Val Pro Lys Pro Cys Cys Ala Pro Thr Gln 385 390 395 400

Leu Asn Ala Ile Ser Val Leu Tyr Phe Asp Asp Ser Ser Asn Val Ile 405 410 415

Leu Lys Lys Tyr Arg Asn Met Val Val Arg Ala Cys Gly Cys His 420 425 430

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<212> PRT

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Lys Trp Ile His Glu Pro Lys Gly Tyr His Ala Asn Phe Cys Leu Gly
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Pro Cys Pro Tyr Ile Trp Ser Leu Asp Thr Gln Tyr Ser Lys Val Leu 35 40 45

Ala Leu Tyr Asn Gln His Asn Pro Gly Ala Ser Ala Ala Pro Cys Cys 50 55 60

Val Pro Gln Ala Leu Glu Pro Leu Pro Ile Val Tyr Tyr Val Gly Arg
65 70 75 80

Lys Pro Lys Val Glu Gln Leu Ser Asn Met Ile Val Arg Ser Cys Lys 85 90 95

Cys Ser

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<213> Homo sapiens
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Lys Trp Ile His Glu Pro Lys Gly Tyr Asn Ala Asn Phe Cys Ala Gly
Ala Cys Pro Tyr Leu Trp Ser Ser Asp Thr Gln His Ser Arg Val Leu
                             40
Ser Leu Tyr Asn Thr Ile Asn Pro Glu Ala Ser Ala Ser Pro Cys Cys
Val Ser Gln Asp Leu Glu Pro Leu Thr Ile Leu Tyr Tyr Ile Gly Lys
Thr Pro Lys Ile Glu Gln Leu Ser Asn Met Ile Val Lys Ser Cys Lys
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Cys Ser
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<400> 42
Cys Cys Val Arg Pro Leu Tyr Ile Asp Phe Arg Gln Asp Leu Gly Trp
Lys Trp Val His Glu Pro Lys Gly Tyr Tyr Ala Asn Phe Cys Ser Gly
                                  25
Pro Cys Pro Tyr Leu Arg Ser Ala Asp Thr Thr His Ser Thr Val Leu
         35
Gly Leu Tyr Asn Thr Leu Asn Pro Glu Ala Ser Ala Ser Pro Cys Cys
Val Pro Gln Asp Leu Glu Pro Leu Thr Ile Leu Tyr Tyr Val Gly Arg
                                          75
Thr Pro Lys Val Glu Gln Leu Ser Asn Met Val Val Lys Ser Cys Lys
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85 90 95

Cys Ser

<210> 43

<211> 98

<212> PRT

<213> Gallus gallus

<220>

<223> TGF-Beta4

<400> 43

Cys Cys Val Arg Pro Leu Tyr Ile Asp Phe Arg Lys Asp Leu Gln Trp

1 5 10 15

Lys Trp Ile His Glu Pro Lys Gly Tyr Met Ala Asn Phe Cys Met Gly
20 25 30

Pro Cys Pro Tyr Ile Trp Ser Ala Asp Thr Gln Tyr Thr Lys Val Leu 35 40 45

Ala Leu Tyr Asn Gln His Asn Pro Gly Ala Ser Ala Ala Pro Cys Cys 50 55 60

Val Pro Gln Thr Leu Asp Pro Leu Pro Ile Ile Tyr Tyr Val Gly Arg
65 70 75 80

Asn Val Arg Val Glu Gln Leu Ser Asn Met Val Val Arg Ala Cys Lys 85 90 95

Cys Ser

<210> 44

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<212> PRT

<213> Xenopus laevis

<220>

<223> TGF-Beta5

<400> 44

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1 5 10 15

Lys Trp Ile His Glu Pro Lys Gly Tyr Glu Ala Asn Tyr Cys Leu Gly
20 25 30

Asn Cys Pro Tyr Ile Trp Ser Met Asp Thr Gln Tyr Ser Lys Val Leu 35 40 45

Ser Leu Tyr Asn Gln Asn Asn Pro Gly Ala Ser Ile Ser Pro Cys Cys 50 55 60

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Val Pro Asp Val Leu Glu Pro Leu Pro Ile Ile Tyr Tyr Val Gly Arg
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Thr Ala Lys Val Glu Gln Leu Ser Asn Met Val Val Arg Ser Cys Asn
Cys Ser
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<213> Drosophila melanogaster
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Lys Cys Pro Phe Pro Leu Ala Asp His Phe Asn Ser Thr Asn His Ala
Val Val Gln Thr Leu Val Asn Asn Met Asn Pro Gly Lys Val Pro Lys
Ala Cys Cys Val Pro Thr Gln Leu Asp Ser Val Ala Met Leu Tyr Leu
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                                  25
             20
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Glu Cys Pro Tyr Pro Leu Thr Glu Ile Leu Asn Gly Ser Asn His Ala 35 40 45

Ile Leu Gln Thr Leu Val His Ser Ile Glu Pro Glu Asp Ile Pro Leu 50 55 60

Pro Cys Cys Val Pro Thr Lys Met Ser Pro Ile Ser Met Leu Phe Tyr 65 70 75 80

Asp Asn Asn Asp Asn Val Val Leu Arg His Tyr Glu Asn Met Ala Val 85 90 95

Asp Glu Cys Gly Cys Arg 100

<210> 47

<211> 102

<212> PRT

<213> Mus musculus

<220>

<223> VGR1

<400> 47

Cys Lys Lys His Glu Leu Tyr Val Ser Phe Gln Asp Leu Gly Trp Gln 1 5 10 15

Asp Trp Ile Ile Ala Pro Lys Gly Tyr Ala Ala Asn Tyr Cys Asp Gly 20 25 30

Glu Cys Ser Phe Pro Leu Asn Ala His Met Asn Ala Thr Asn His Ala 35 40 45

Ile Val Gln Thr Leu Val His Leu Met Asn Pro Glu Tyr Val Pro Lys 50 55 60

Pro Cys Cys Ala Pro Thr Lys Leu Asn Ala Ile Ser Val Leu Tyr Phe 65 70 75 80

Asp Asp Asn Ser Asn Val Ile Leu Lys Lys Tyr Arg Asn Met Val Val 85 90 95

Arg Ala Cys Gly Cys His 100

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<211> 118

<212> PRT

<213> Drosophila melanogaster

<220>

<223> 60A

<400> 48

Cys Gln Met Gln Thr Leu Tyr Ile Asp Phe Lys Asp Leu Gly Trp His

Asp Trp Ile Ile Ala Pro Glu Gly Tyr Gly Ala Phe Tyr Cys Ser Gly

Glu Cys Asn Phe Pro Leu Asn Ala His Met Asn Ala Thr Asn His Ala

Ile Val Gln Thr Leu Val His Leu Leu Glu Pro Lys Lys Val Pro Lys 55

Pro Cys Cys Ala Pro Thr Arg Leu Gly Ala Leu Pro Val Leu Tyr His

Pro Cys Cys Ala Pro Thr Arg Leu Gly Ala Leu Pro Val Leu Tyr His

Leu Asn Asp Glu Asn Val Asn Leu Lys Lys Tyr Arg Asn Met Ile Val

Lys Ser Cys Gly Cys His 115

<210> 49

<211> 101

<212> PRT

<213> Homo sapiens

<220>

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<223> BMP-2A

<400> 49

Cys Lys Arg His Pro Leu Tyr Val Asp Phe Ser Asp Val Gly Trp Asn

Asp Trp Ile Val Ala Pro Pro Gly Tyr His Ala Phe Tyr Cys His Gly

Glu Cys Pro Phe Pro Leu Ala Asp His Leu Asn Ser Thr Asn His Ala

Ile Val Gln Thr Leu Val Asn Ser Val Asn Ser Lys Ile Pro Lys Ala 55

Cys Cys Val Pro Thr Glu Leu Ser Ala Ile Ser Met Leu Tyr Leu Asp

Glu Asn Glu Lys Val Val Leu Lys Asn Tyr Gln Asp Met Val Val Glu 90

Gly Cys Gly Cys Arg 100

<210> 50

<211> 103

Gly Cys Gly Cys Arg

```
<212> PRT
<213> Homo sapiens
<220>
<223> BMP3
<400> 50
Cys Ala Arg Arg Tyr Leu Lys Val Asp Phe Ala Asp Ile Gly Trp Ser
Glu Trp Ile Ile Ser Pro Lys Ser Phe Asp Ala Tyr Tyr Cys Ser Gly
Ala Cys Gln Phe Pro Met Pro Lys Ser Leu Lys Pro Ser Asn His Ala
Thr Ile Gln Ser Ile Val Arg Ala Val Gly Val Val Pro Gly Ile Pro
Glu Pro Cys Cys Val Pro Glu Lys Met Ser Ser Leu Ser Ile Leu Phe
Phe Asp Glu Asn Lys Asn Val Val Leu Lys Val Tyr Pro Asn Met Thr
Val Glu Ser Cys Ala Cys Arg
            100
<210> 51
<211> 101
<212> PRT
<213> Homo sapiens
<220>
<223> BMP-4
<400> 51
Cys Arg Arg His Ser Leu Tyr Val Asp Phe Ser Asp Val Gly Trp Asn
Asp Trp Ile Val Ala Pro Pro Gly Tyr Gln Ala Phe Tyr Cys His Gly
Asp Cys Pro Phe Pro Leu Ala Asp His Leu Asn Ser Thr Asn His Ala
Ile Val Gln Thr Leu Val Asn Ser Val Asn Ser Ser Ile Pro Lys Ala
Cys Cys Val Pro Thr Glu Leu Ser Ala Ile Ser Met Leu Tyr Leu Asp
Glu Tyr Asp Lys Val Val Leu Lys Asn Tyr Gln Glu Met Val Val Glu
```

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<210> 52
<211> 102
<212> PRT
<213> Homo sapiens
<220>
<223> BMP-5
<400> 52
Cys Lys Lys His Glu Leu Tyr Val Ser Phe Arg Asp Leu Gly Trp Gln
Asp Trp Ile Ile Ala Pro Glu Gly Tyr Ala Ala Phe Tyr Cys Asp Gly
                                 25
             20
Glu Cys Ser Phe Pro Leu Asn Ala His Met Asn Ala Thr Asn His Ala
Ile Val Gln Thr Leu Val His Leu Met Phe Pro Asp His Val Pro Lys
Pro Cys Cys Ala Pro Thr Lys Leu Asn Ala Ile Ser Val Leu Tyr Phe
                     70
Asp Asp Ser Ser Asn Val Ile Leu Lys Lys Tyr Arg Asn Met Val Val
Arg Ser Cys Gly Cys His
            100
<210> 53
<211> 102
<212> PRT
<213> Homo sapiens
<220>
<223> BMP-6
<400> 53
Cys Arg Lys His Glu Leu Tyr Val Ser Phe Gln Asp Leu Gly Trp Gln
Asp Trp Ile Ile Ala Pro Lys Gly Tyr Ala Ala Asn Tyr Cys Asp Gly
Glu Cys Ser Phe Pro Leu Asn Ala His Met Asn Ala Thr Asn His Ala
                                                  45
         35
Ile Val Gln Thr Leu Val His Leu Met Asn Pro Glu Tyr Val Pro Lys
Pro Cys Cys Ala Pro Thr Lys Leu Asn Ala Ile Ser Val Leu Tyr Phe
```

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Asp Asp Asn Ser Asn Val Ile Leu Lys Lys Tyr Arg Asn Met Val Val
                                     90
Arg Ala Cys Gly Cys His
            100
<210> 54
<211> 103
<212> PRT
<213> Gallus gallus
<220>
<223> DORSALIN
<400> 54
Cys Arg Arg Thr Ser Leu His Val Asn Phe Lys Glu Ile Gly Trp Asp
                  5
Ser Trp Ile Ile Ala Pro Lys Asp Tyr Glu Ala Phe Glu Cys Lys Gly
Gly Cys Phe Phe Pro Leu Thr Asp Asn Val Thr Pro Thr Lys His Ala
Ile Val Gln Thr Leu Val His Leu Gln Asn Pro Lys Lys Ala Ser Lys
    50
Ala Cys Cys Val Pro Thr Lys Leu Asp Ala Ile Ser Ile Leu Tyr Lys
Asp Asp Ala Gly Val Pro Thr Leu Ile Tyr Asn Tyr Glu Gly Met Lys
Val Ala Glu Cys Gly Cys Arg
            100
<210> 55
<211> 102
<212> PRT
<213> Homo sapiens
<220>
<223> OP-1
<400> 55
Cys Lys Lys His Glu Leu Tyr Val Ser Phe Arg Asp Leu Gly Trp Gln
Asp Trp Ile Ile Ala Pro Glu Gly Tyr Ala Ala Tyr Tyr Cys Glu Gly
```

Glu Cys Ala Phe Pro Leu Asn Ser Tyr Met Asn Ala Thr Asn His Ala

```
Ile Val Gln Thr Leu Val His Phe Ile Asn Pro Glu Thr Val Pro Lys 50 55 60
```

Pro Cys Cys Ala Pro Thr Gln Leu Asn Ala Ile Ser Val Leu Tyr Phe 65 70 75 80

Asp Asp Ser Ser Asn Val Ile Leu Lys Lys Tyr Arg Asn Met Val Val 85 90 95

Arg Ala Cys Gly Cys His 100

<210> 56

<211> 102

<212> PRT

<213> Homo sapiens

<220>

<223> OP-2

<400> 56

Cys Arg Arg His Glu Leu Tyr Val Ser Phe Gln Asp Leu Gly Trp Leu 1 5 10 15

Asp Trp Val Ile Ala Pro Gln Gly Tyr Ser Ala Tyr Tyr Cys Glu Gly
20 25 30

Glu Cys Ser Phe Pro Leu Asp Ser Cys Met Asn Ala Thr Asn His Ala 35 40 45

Ile Leu Gln Ser Leu Val His Leu Met Lys Pro Asn Ala Val Pro Lys 50 55 60

Ala Cys Cys Ala Pro Thr Lys Leu Ser Ala Thr Ser Val Leu Tyr Tyr 65 70 75 80

Asp Ser Ser Asn Asn Val Ile Leu Arg Lys His Arg Asn Met Val Val 85 90 95

Lys Ala Cys Gly Cys His 100

<210> 57

<211> 102

<212> PRT

<213> Mus musculus

<220>

<223> OP-3

<400> 57

Cys Arg Arg His Glu Leu Tyr Val Ser Phe Arg Asp Leu Gly Trp Leu $1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15$

Asp Ser Val Ile Ala Pro Gln Gly Tyr Ser Ala Tyr Tyr Cys Ala Gly

20 25 30

Glu Cys Ile Tyr Pro Leu Asn Ser Cys Met Asn Ser Thr Asn His Ala 35 40 45

Thr Met Gln Ala Leu Val His Leu Met Lys Pro Asp Ile Ile Pro Lys 50 55 60

Val Cys Cys Val Pro Thr Glu Leu Ser Ala Ile Ser Leu Leu Tyr Tyr 65 70 75 80

Asp Arg Asn Asn Val Ile Leu Arg Arg Glu Arg Asn Met Val Val 85 90 95

Gln Ala Cys Gly Cys His 100

<210> 58

<211> 107

<212> PRT

<213> Mus musculus

<220>

<223> GDF-1

<400> 58

Cys Arg Thr Arg Arg Leu His Val Ser Phe Arg Glu Val Gly Trp His 1 5 10 15

Arg Trp Val Ile Ala Pro Arg Gly Phe Leu Ala Asn Phe Cys Gln Gly
20 25 30

Thr Cys Ala Leu Pro Glu Thr Leu Arg Gly Pro Gly Gly Pro Pro Ala 35 40 45

Leu Asn His Ala Val Leu Arg Ala Leu Met His Ala Ala Pro Thr 50 55 60

Pro Gly Ala Gly Ser Pro Cys Cys Val Pro Glu Arg Leu Ser Pro Ile 65 70 75 80

Ser Val Leu Phe Phe Asp Asn Ser Asp Asn Val Val Leu Arg His Tyr 85 90 95

Glu Asp Met Val Val Asp Glu Cys Gly Cys Arg 100 105

<210> 59

<211> 101

<212> PRT

<213> Mus musculus

<220>

<223> GDF-3

<400> 59 Cys His Arg His Gln Leu Phe Ile Asn Phe Gln Asp Leu Gly Trp His Lys Trp Val Ile Ala Pro Lys Gly Phe Met Ala Asn Tyr Cys His Gly Glu Cys Pro Phe Ser Met Thr Thr Tyr Leu Asn Ser Ser Asn Tyr Ala 40 Phe Met Gln Ala Leu Met His Met Ala Asp Pro Lys Val Pro Lys Ala 55 Val Cys Val Pro Thr Lys Leu Ser Pro Ile Ser Met Leu Tyr Gln Asp Ser Asp Lys Asn Val Ile Leu Arg His Tyr Glu Asp Met Val Val Asp 90 Glu Cys Gly Cys Gly 100 <210> 60 <211> 102 <212> PRT <213> Mus musculus <220> <223> GDF-9 <400> 60 Cys Glu Leu His Asp Phe Arg Leu Ser Phe Ser Gln Leu Lys Trp Asp 15 5 Asn Trp Ile Val Ala Pro His Arg Tyr Asn Pro Arg Tyr Cys Lys Gly 25 Asp Cys Pro Arg Ala Val Arg His Arg Tyr Gly Ser Pro Val His Thr 40

Met Val Gln Asn Ile Ile Tyr Glu Lys Leu Asp Pro Ser Val Pro Arg
50 55 60

Pro Ser Cys Val Pro Gly Lys Tyr Ser Pro Leu Ser Val Leu Thr Ile
65 70 75 80

Glu Pro Asp Gly Ser Ile Ala Tyr Lys Glu Tyr Glu Asp Met Ile Ala 85 90 95

Thr Arg Cys Thr Cys Arg 100

<210> 61 <211> 105 <212> PRT

```
<213> Homo sapiens
<220>
<223> INHIBIN-Alpha
<400> 61
Cys His Arg Val Ala Leu Asn Ile Ser Phe Gln Glu Leu Gly Trp Glu
Arg Trp Ile Val Tyr Pro Pro Ser Phe Ile Phe His Tyr Cys His Gly
                                 25
Gly Cys Gly Leu His Ile Pro Pro Asn Leu Ser Leu Pro Val Pro Gly
                             40
Ala Pro Pro Thr Pro Ala Gln Pro Tyr Ser Leu Leu Pro Gly Ala Gln
Pro Cys Cys Ala Ala Leu Pro Gly Thr Met Arg Pro Leu His Val Arg_
Thr Thr Ser Asp Gly Gly Tyr Ser Phe Lys Tyr Glu Thr Val Pro Asn
                                90
Leu Leu Thr Gln His Cys Ala Cys Ile
            100
<210> 62
<211> 106
<212> PRT
<213> Bos taurus
<220>
<223> INHIBIN-BetaA
<400> 62
Cys Cys Lys Lys Gln Phe Phe Val Ser Phe Lys Asp Ile Gly Trp Asn
Asp Trp Ile Ile Ala Pro Ser Gly Tyr His Ala Asn Tyr Cys Glu Gly
Glu Cys Pro Ser His Ile Ala Gly Thr Ser Gly Ser Ser Leu Ser Phe
His Ser Thr Val Ile Asn His Tyr Arg Met Arg Gly His Ser Pro Phe
Ala Asn Leu Lys Ser Cys Cys Val Pro Thr Lys Leu Arg Pro Met Ser
Met Leu Tyr Tyr Asp Asp Gly Gln Asn Ile Ile Lys Lys Asp Ile Gln
```

Asn Met Ile Val Glu Glu Cys Gly Cys Ser

```
<210> 63
<211> 106
<212> PRT
<213> Homo sapiens
<220>
<223> INHIBIN-BetaB
<400> 63
Cys Cys Lys Lys Gln Phe Phe Val Ser Phe Lys Asp Ile Gly Trp Asn
Asp Trp Ile Ile Ala Pro Ser Gly Tyr His Ala Asn Tyr Cys Glu Gly
Glu Cys Pro Ser His Ile Ala Gly Thr Ser Gly Ser Ser Leu Ser Phe
His Ser Thr Val Ile Asn His Tyr Arg Met Arg Gly His Ser Pro Phe
Ala Asn Leu Lys Ser Cys Cys Val Pro Thr Lys Leu Arg Pro Met Ser
Met Leu Tyr Tyr Asp Asp Gly Gln Asn Ile Ile Lys Lys Asp Ile Gln
Asn Met Ile Val Glu Glu Cys Gly Cys Ser
            100
<210> 64
<211> 98
<212> PRT
<213> Artificial Sequence
<223> Description of Artificial Sequence: TGF-B
      SUBGROUP SEQUENCE PATTERN
<220>
<223> Each Xaa is independently selected from a group of
      one or more specified amino acids as defined in
      the specification
<400> 64
Cys Cys Val Arg Pro Leu Tyr Ile Asp Phe Arg Xaa Asp Leu Gly Trp
Lys Trp Ile His Glu Pro Lys Gly Tyr Xaa Ala Asn Phe Cys Xaa Gly
Xaa Cys Pro Tyr Xaa Trp Ser Xaa Asp Thr Gln Xaa Ser Xaa Val Leu
```

```
Xaa Leu Tyr Asn Xaa Xaa Asn Pro Xaa Ala Ser Ala Xaa Pro Cys Cys 50 55 60
```

Val Pro Gln Xaa Leu Glu Pro Leu Xaa Ile Xaa Tyr Tyr Val Gly Arg 65 70 75 80

Xaa Xaa Lys Val Glu Gln Leu Ser Asn Met Xaa Val Xaa Ser Cys Lys 85 90 95

Cys Ser

<210> 65

<211> 104

<212> PRT

<213> Artificial Sequence

<220>

<223> Each Xaa is independently selected from a group of one or more specified amino acids as defined in the specification

<220>

<223> Description of Artificial Sequence: VG/DPP
SUBGROUP SEQUENCE PATTERN

<400> 65

Cys Xaa Xaa Xaa Leu Tyr Val Xaa Phe Xaa Asp Xaa Gly Trp Xaa 1 5 10 15

Asp Trp Ile Ile Ala Pro Xaa Gly Tyr Xaa Ala Xaa Tyr Cys Xaa Gly
20 25 30

Xaa Cys Xaa Phe Pro Leu Xaa Xaa Xaa Xaa Asn Xaa Thr Asn His Ala 35 40 45

Ile Xaa Gln Thr Leu Val Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Aaa Pro 50 55 60

Lys Xaa Cys Cys Xaa Pro Thr Xaa Leu Xaa Ala Xaa Ser Xaa Leu Tyr 65 70 75 80

Xaa Asp Xaa Xaa Xaa Xaa Val Xaa Leu Xaa Xaa Tyr Xaa Xaa Met 85 90 95

Xaa Val Xaa Xaa Cys Gly Cys Xaa 100

<210> 66

<211> 107

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: GDF SUBGROUP

PATTERN

<220>

<223> Each Xaa is independently selected from a group of one or more specified amino acids as defined in the specification

<400> 66

Cys Xaa Xaa Xaa Xaa Xaa Xaa Xaa Phe Xaa Xaa Xaa Trp Xaa 1 10 15

Xaa Trp Xaa Xaa Ala Pro Xaa Xaa Xaa Xaa Xaa Xaa Cys Xaa Gly
20 25 30

Pro Xaa Xaa Xaa Xaa Xaa Cys Val Pro Xaa Xaa Ser Pro Xaa 65 70 75 80

Glu Asp Met Xaa Xaa Xaa Xaa Cys Xaa Cys Xaa 100 105

<210> 67

<211> 109

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: INHIBIN
SUBGROUP PATTERN

<220>

<223> Each Xaa is independently selected from a group of one or more specified amino acids as defined in the specification

<400> 67

Cys Xaa Xaa Xaa Xaa Xaa Xaa Xaa Phe Xaa Xaa Gly Trp Xaa 1 5 10 15

Xaa Trp Ile Xaa Xaa Pro Xaa Xaa Xaa Xaa Xaa Xaa Tyr Cys Xaa Gly
20 25 30

Xaa Xaa Xaa Xaa Xaa Cys Cys Xaa Xaa Xaa Pro Xaa Xaa Xaa Xaa Xaa 65
70
75
80

Xaa Xaa Xaa Asn Xaa Xaa Xaa Xaa Cys Xaa Cys Xaa 100 105

<210> 68

<211> 139

<212> PRT

<213> Homo sapiens

<220>

<223> Mature H2223 mutant

<400> 68

Ser Thr Gly Ser Lys Gln Arg Ser Gln Asn Arg Ser Lys Thr Pro Lys

1 5 10 15

Asn Gln Glu Ala Leu Arg Met Ala Asn Val Ala Glu Asn Ser Ser Ser 20 25 30

Asp Gln Arg Gln Ala Cys Lys Lys His Glu Leu Tyr Val Ser Phe Arg 35 40 45

Asp Leu Gly Trp Gln Asp Trp Ile Ile Ala Pro Glu Gly Tyr Ala Ala 50 55 60

Tyr Tyr Cys Glu Gly Glu Cys Ala Phe Pro Leu Asn Ser Tyr Met Asn 65 70 75 80

Ala Thr Asn His Ala Ile Val Gln Thr Leu Val His Phe Ile Asn Pro 85 90 95

Glu Thr Val Pro Lys Pro Cys Cys Ala Pro Thr Gln Leu Asn Ala Ile 100 105 110

Ser Val Leu Tyr Phe Asp Asp Ser Ser Asn Val Ile Leu Lys Lys Tyr 115 120 125

Glu Asp Met Val Val Glu Ala Cys Gly Cys Arg 130 135

<210> 69

<211> 117

<212> PRT

<213> Homo sapiens

<220>

<223> Trypsin truncated H2223 mutant

<400> 69

Met Ala Asn Val Ala Glu Asn Ser Ser Ser Asp Gln Arg Gln Ala Cys

```
Lys Lys His Glu Leu Tyr Val Ser Phe Arg Asp Leu Gly Trp Gln Asp
             20 -
                                 25
Trp Ile Ile Ala Pro Glu Gly Tyr Ala Ala Tyr Tyr Cys Glu Gly Glu
Cys Ala Phe Pro Leu Asn Ser Tyr Met Asn Ala Thr Asn His Ala Ile
                         55
Val Gln Thr Leu Val His Phe Ile Asn Pro Glu Thr Val Pro Lys Pro
Cys Cys Ala Pro Thr Gln Leu Asn Ala Ile Ser Val Leu Tyr Phe Asp
Asp Ser Ser Asn Val Ile Leu Lys Lys Tyr Glu Asp Met Val Val Glu
                                105
Ala Cys Gly Cys Arg
        115
<210> 70
<211> 33
<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence: Primer #1
<220>
<221> CDS
<222> (1)..(33)
<400> 70
                                                                   33
gcg ccc acg cag ctc agc gct atc tcc gtc ctc
Ala Pro Thr Gln Leu Ser Ala Ile Ser Val Leu
<210> 71
<211> 11
<212> PRT
<213> Artificial Sequence
Ala Pro Thr Gln Leu Ser Ala Ile Ser Val Leu
                  5
1
<210> 72
<211> 43
<212> DNA
<213> Artificial Sequence
```

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<220>
     <223> Description of Artificial Sequence: Primer #2
     <400> 72
                                                                          43
     ctatctgcag ccacaagett cgaccaccat gtcttcgtat ttc
     <210> 73
     <211> 43
     <212> DNA
     <213> Artificial Sequence
     <220>
     <223> Description of Artificial Sequence:complement of
           Primer #2
     <220>
     <221> CDS
     <222> (2)..(43)
     <400> 73
                                                                          43
     g aaa tac gaa gac atg gtg gtc gaa gct tgt ggc tgc aga tag
       Lys Tyr Glu Asp Met Val Val Glu Ala Cys Gly Cys Arg
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     <210> 74
<211> 13
     <212> PRT
IJ
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     <213> Artificial Sequence
1,,3
     <400> 74
12
     Lys Tyr Glu Asp Met Val Val Glu Ala Cys Gly Cys Arg
                                            10
                        5
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<210> 75
     <211> 44
     <212> DNA
     <213> Artificial Sequence
     <220>
     <223> Description of Artificial Sequence: the sequence
           between the T7 promoter, at the XbaI site, and the
           ATG codon
     <400> 75
                                                                          44
     tctagaataa ttttgtttaa cctttaagaa ggagatatac gatg
     <210> 76
     <211> 19
     <212> DNA
     <213> Artificial Sequence
     <220>
     <223> Description of Artificial Sequence: Primer #3
```

```
<400> 76
                                                                            19
      taatacgact cactatagg
      <210> 77
     <211> 18
      <212> DNA
     <213> Artificial Sequence
      <223> Description of Artificial Sequence: Primer #4
     <400> 77
                                                                            18
      gctgagctgc gtgggcgc
     <210> 78
      <211> 18
      <212> DNA
      <213> Artificial Sequence
     <220>
      <223> Description of Artificial Sequence: complement of
: []
            Primer #4
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·+, }
      <220>
<221> CDS
     <222> (1)..(18)
اً إِنْ إِنَّا
١,,]
     <400> 78
إإبإ
                                                                            18
     gcg ccc acg cag ctc agc
15
     Ala Pro Thr Gln Leu Ser
                         5
        1
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-11
<210> 79
     <211> 6
<212> PRT
<213> Artificial Sequence
      <400> 79
      Ala Pro Thr Gln Leu Ser
      <210> 80
      <211> 23
      <212> DNA
      <213> Artificial Sequence
      <223> Description of Artificial Sequence: Primer #5
      <400> 80
                                                                            23
      ggatcctatc tgcagccaca agc
```

```
<210> 81
     <211> 23
     <212> DNA
     <213> Artificial Sequence
     <223> Description of Artificial Sequence:, complement of
           Primer #5
     <220>
     <221> CDS
     <222> (1)..(18)
     <400> 81
                                                                          23
     gct tgt ggc tgc aga tag gatcc
     Ala Cys Gly Cys Arg
     <210> 82
     <211> 5
     <212> PRT
     <213> Artificial Sequence
4.
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     <400> 82
١٠, إ
     Ala Cys Gly Cys Arg
IJ,Ĵ
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ij
     <210> 83
     <211> 102
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     <212> PRT
     <213> Homo sapiens
en k
     <220>
     <223> CDMP-1/GDF-5
1,11
     <400> 83
Ų
     Cys Ser Arg Lys Ala Leu His Val Asn Phe Lys Asp Met Gly Trp Asp
                                            10
     Asp Trp Ile Ile Ala Pro Leu Glu Tyr Glu Ala Phe His Cys Glu Gly
     Leu Cys Glu Phe Pro Leu Arg Ser His Leu Glu Pro Thr Asn His Ala
              35
     Val Ile Gln Thr Leu Met Asn Ser Met Asp Pro Glu Ser Thr Pro Pro
     Thr Cys Cys Val Pro Thr Arg Leu Ser Pro Ile Ser Ile Leu Phe Ile
     Asp Ser Ala Asn Asn Val Val Tyr Lys Gln Tyr Glu Asp Met Val Val
```

Glu Ser Cys Gly Cys Arg 100

<210> 84

<211> 102

<212> PRT

<213> Homo sapiens

<220>

<223> CDMP-2/GDF-6

<400> 84

Cys Ser Lys Lys Pro Leu His Val Asn Phe Lys Glu Leu Gly Trp Asp 1 5 10 15

Asp Trp Ile Ile Ala Pro Leu Glu Tyr Glu Ala Tyr His Cys Glu Gly
20 25 30

Val Cys Asp Phe Pro Leu Arg Ser His Leu Glu Pro Thr Asn His Ala 35 40 45

Ile Ile Gln Thr Leu Met Asn Ser Met Asp Pro Gly Ser Thr Pro Pro 50 55 60

Ser Cys Cys Val Pro Thr Lys Leu Thr Pro Ile Ser Ile Leu Tyr Ile 65 70 75 80

Asp Ala Gly Asn Asn Val Val Tyr Lys Gln Tyr Glu Asp Met Val Val 85 90 95

Glu Ser Cys Gly Cys Arg 100

<210> 85

<211> 102

<212> PRT

<213> Mus musculus

<220>

<223> GDF-6

<400> 85

Cys Ser Arg Lys Pro Leu His Val Asn Phe Lys Glu Leu Gly Trp Asp 1 5 10 15

Asp Trp Ile Ile Ala Pro Leu Glu Tyr Glu Ala Tyr His Cys Glu Gly 20 25 30

Val Cys Asp Phe Pro Leu Arg Ser His Leu Glu Pro Thr Asn His Ala 35 40 45

Ile Ile Gln Thr Leu Met Asn Ser Met Asp Pro Gly Ser Thr Pro Pro 50 55 60

Ser Cys Cys Val Pro Thr Lys Leu Thr Pro Ile Ser Ile Leu Tyr Ile

65 70 75 80

Asp Ala Gly Asn Asn Val Val Tyr Lys Gln Tyr Glu Asp Met Val Val 85 90 95

Glu Ser Cys Gly Cys Arg 100

<210> 86

<211> 102

<212> PRT

<213> Bos taurus

<220>

<223> CDMP-2

<400> 86

Cys Ser Lys Lys Pro Leu His Val Asn Phe Lys Glu Leu Gly Trp Asp 1 5 10 15

Asp Trp Ile Ile Ala Pro Leu Glu Tyr Glu Ala Tyr His Cys Glu Gly
20 25 30

Val Cys Asp Phe Pro Leu Arg Ser His Leu Glu Pro Thr Asn His Ala 35 40 45

Ile Ile Gln Thr Leu Met Asn Ser Met Asp Pro Gly Ser Thr Pro Pro 50 55 60

Ser Cys Cys Val Pro Thr Lys Leu Thr Pro Ile Ser Ile Leu Tyr Ile 65 70 75 80

Asp Ala Gly Asn Asn Val Val Tyr Asn Glu Tyr Glu Glu Met Val Val 85 90 95

Glu Ser Cys Gly Cys Arg 100

<210> 87

<211> 102

<212> PRT

<213> Mus musculus

<220>

<223> GDF-7

<400> 87

Cys Ser Arg Lys Ser Leu His Val Asp Phe Lys Glu Leu Gly Trp Asp 1 5 10 15

Asp Trp Ile Ile Ala Pro Leu Asp Tyr Glu Ala Tyr His Cys Glu Gly
20 25 30

Val Cys Asp Phe Pro Leu Arg Ser His Leu Glu Pro Thr Asn His Ala 35 40 45

<210> 88 <211> 102 <212> PRT <213> Homo sapiens <220> <223> CDMP-3 construct

Asp Trp Ile Ile Ala Pro Leu Asp Tyr Glu Ala Tyr His Cys Glu Gly
20 . 30

Leu Cys Asp Phe Pro Leu Arg Ser His Leu Glu Pro Thr Asn His Ala
35 40 45

Ile Ile Gln Thr Leu Leu Asn Ser Met Ala Pro Asp Ala Ala Pro Ala 50 55 60

Ser Cys Cys Val Pro Ala Arg Leu Ser Pro Ile Ser Ile Leu Tyr Ile
65 70 75 80

Asp Ala Ala Asn Asn Val Val Tyr Lys Glh Tyr Glu Asp Met Val Val 85 90 95

Glu Ala Cys Gly Cys Arg 100